Inhibitors of Memapsin 2 and Use Thereof ld Koelsch, Jordan J. N. Tang, Lin Hong, and Arun K of U.S.S.N. 09/603,713

April 27, 2001 Atty. Docket No.: OMRF 182 DIV

FIG.

NILVDTGSSN RPDDSLEPFF GIDHSLYTGS GELGTDLVSI DSGTTNLRLP NIFPVISLYL DMEDCGYNIP STGTVMGAVI QHDDFADDIS GKSGQGYYVE MTVGSPPQTL STYRDLRKGV YVPYTQGKWE ILGLAYAEIA NLFSLQLCGA GFPLNQSEVL ASVGGSMIIG INGQDLKMDC KEYNYDKSIV PDGFWLGEQL VCWQAGTTPW PVEDVATSQD DCYKFAISQS ACHVHDEFRT AAVEGPFVTL QWRCLRCLRQ FFINGSNWEG FMLPLCLMVC SFVEMVDNLR LHRYYQRQLS ITILPQQYLR YYEVIIVRVE ARKRIGFAVS NIAAITESDK IKAASSTEKF AYVMAAICAL PHGPNVTVRA DSLVKQTHVP EEPEEPGRRG FAVGAAPHPF LWYTPIRREW KKVFEAAVKS MEGFYVVFDR MGEVTNQSFR QTDESTILMTI LLK* 438 36p 188 288 138 238 338 388 38 8 8

QTDEST GFAVSA The →Promemapsin 2-T2 H→Promemapsin 2-T1 MASMTGGQQMGRGSMAGVL MASMTGGQQMGRGSMAGVL

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LGLRLPRETD

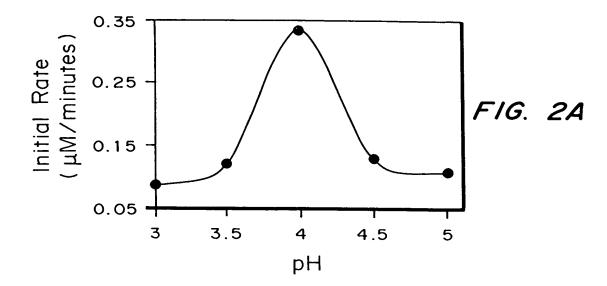
PLRSGLGGAP

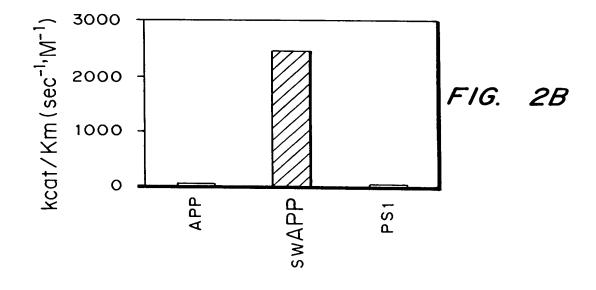
AHGTQHGIRL

1v MASMTGGQQM GRGSMAGVLP

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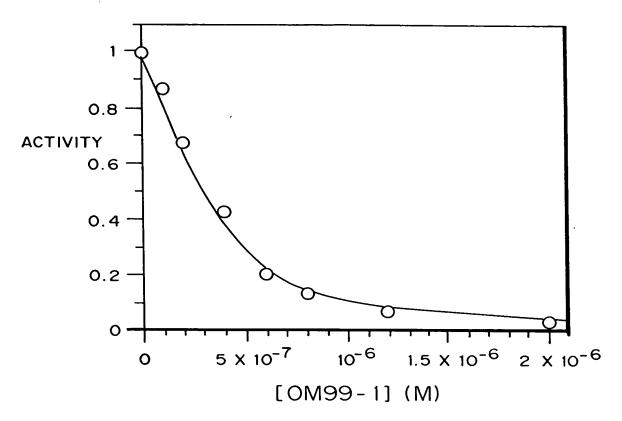
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of U.S.S.N. 09/603,713

Atty. Docket No.: OMRF 182 DIV





[Mep2] = $0.47 \mu M$

DSB4525.O4301

[Fluo. Substrate] = $29.9 \mu M$

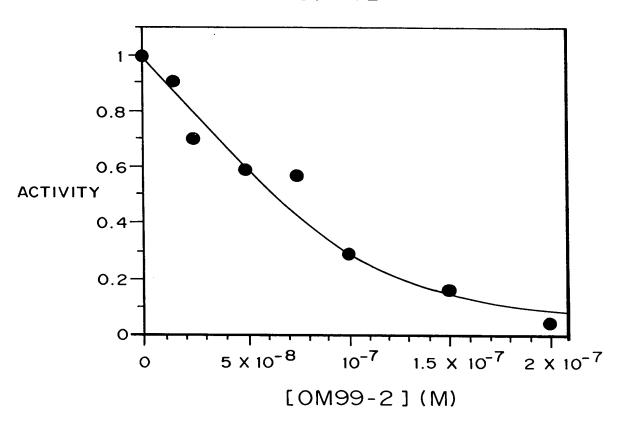
Buffer: Na Acetate 0.1 M, 5% DMSO, pH 4.5 at 37°C

Excitation at 350 nm

Emission at 490 nm

Parameter	Value	Std. Error
Ki	6.84 e - 8	2.72 e - 9

FIG. 4B

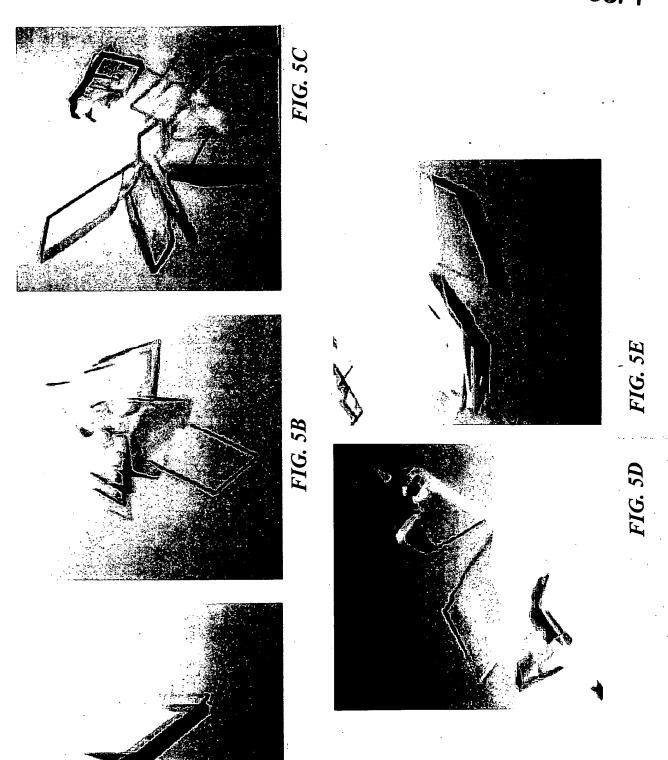


[E]o=O.11 μ M [Fluo Substrate] = $29.9 \mu M$

Parameter	Value	Std. Error
Ki	9.58e-9	2.86e - 10

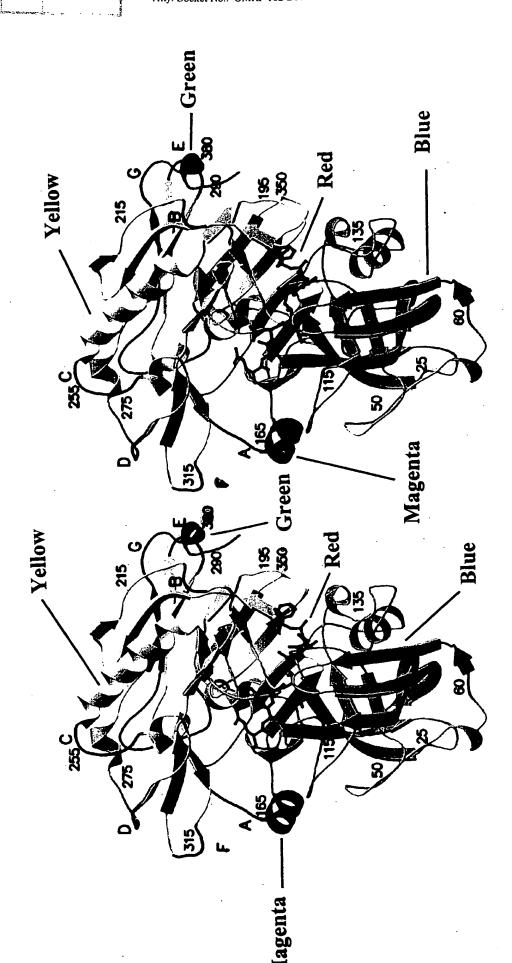
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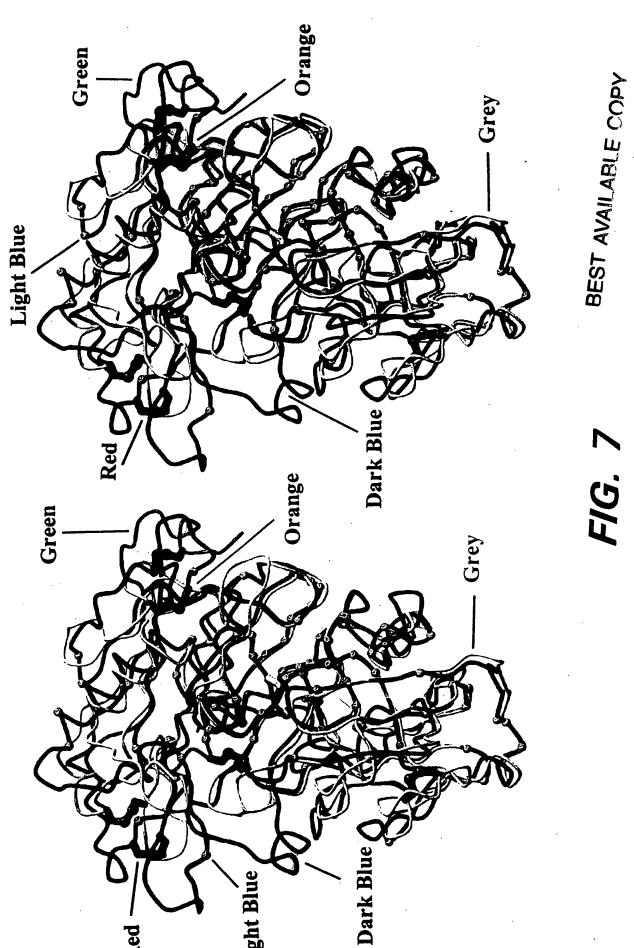


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FIG. 6

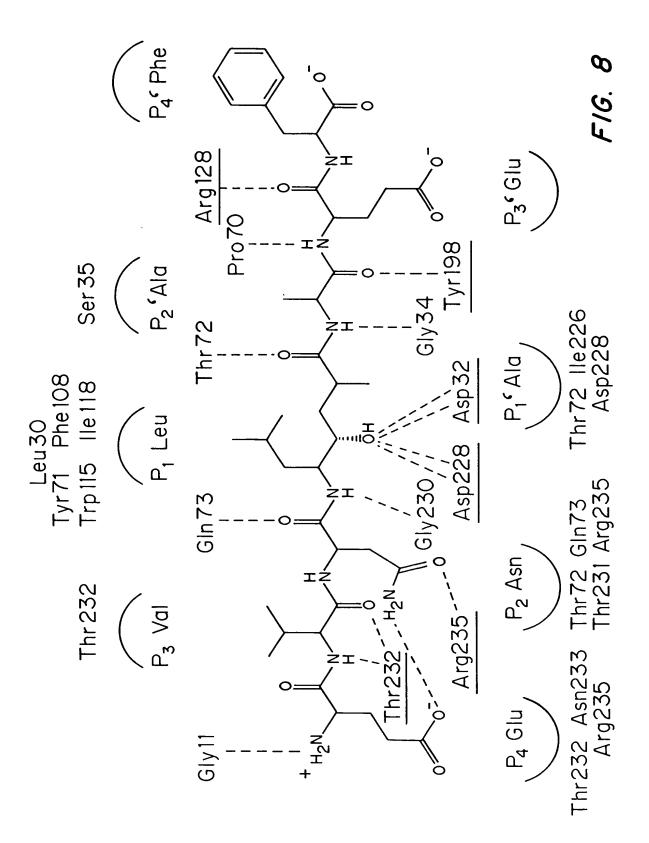


y: Gerald Koelsch, Jordan J. N. Tang, Lin Hong, and Ar. Ghosh ivisional of U.S.S.N. 09/603,713 Filed: April 27, 2001 Atty. Docket No.: OMRF 182 DIV



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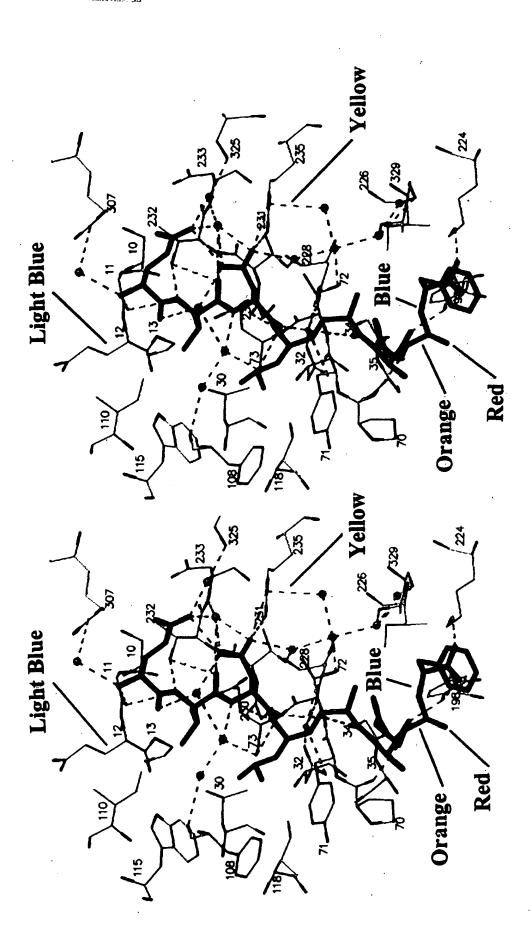
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P₁ Leu P₃ Val H H_2N NH_2 Ē OH R P₁ P₂

FIG. 10

$$P_4$$
 Glu
$$P_2$$
 Asn
$$A \xrightarrow{P_4} H$$

$$P_4 = H$$

FIG. 11

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